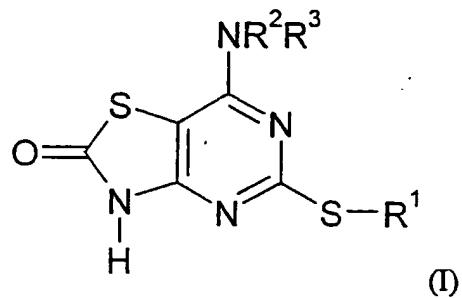


CLAIMS

1. A compound of formula (I) or a pharmaceutically acceptable salt or solvate thereof:



5

in which

R¹ represents a C₃-C₇ carbocyclic, C₁-C₈ alkyl, C₂-C₆ alkenyl or C₂-C₆ alkynyl group, each of the groups being optionally substituted by one or more substituent groups independently selected from halogen atoms, -OR⁴, -NR⁵R⁶, -CONR⁵R⁶, -COOR⁷, -NR⁸COR⁹, -SR¹⁰, -SO₂R¹⁰, -SO₂NR⁵R⁶, -NR⁸SO₂R⁹ or an aryl or heteroaryl group, both of which may be optionally substituted by one or more substituents independently selected from halogen atoms, cyano, nitro, -OR⁴, -NR⁵R⁶, -CONR⁵R⁶, -COOR⁷, -NR⁸COR⁹, -SR¹⁰, -SO₂R¹⁰, -SO₂NR⁵R⁶, -NR⁸SO₂R⁹, C₁-C₆ alkyl or trifluoromethyl groups;

R² and R³ each independently represent a hydrogen atom, or a C₃-C₇ carbocyclic, C₁-C₈ alkyl, C₂-C₆ alkenyl or C₂-C₆ alkynyl group, the latter four groups may be optionally substituted by one or more substituent groups independently selected from:

- (a) halogen atoms, -OR⁴, -NR⁵R⁶, -CONR⁵R⁶, -COOR⁷, -NR⁸COR⁹, -SR¹⁰, -SO₂R¹⁰, -SO₂NR⁵R⁶, -NR⁸SO₂R⁹
- (b) a 3-8 membered ring optionally containing one or more atoms selected from O, S, NR⁸ and itself optionally substituted by C₁-C₃-alkyl or halogen,
- (c) an aryl group or heteroaryl group each of which may be optionally substituted by one or more substituents independently selected from halogen atoms, cyano, nitro, -OR⁴, -NR⁵R⁶, -CONR⁵R⁶, -NR⁸COR⁹, -SO₂NR⁵R⁶, -NR⁸SO₂R⁹, C₁-C₆ alkyl and trifluoromethyl groups;

R⁴ represents hydrogen, C₁-C₆ alkyl or a phenyl group the latter two of which may be optionally substituted by one or more substituent groups independently selected from halogen atoms, phenyl, -OR¹¹ and -NR¹²R¹³

5 R⁵ and R⁶ independently represent a hydrogen atom or a C₁-C₆ alkyl or phenyl group the latter two of which may be optionally substituted by one or more substituent groups independently selected from halogen atoms, phenyl, -OR¹⁴ and -NR¹⁵R¹⁶, -CONR¹⁵R¹⁶, -NR¹⁵COR¹⁶, -SONR¹⁵R¹⁶, NR¹⁵SO₂R¹⁶

or

10 R⁵ and R⁶ together with the nitrogen atom to which they are attached form a 4- to 7-membered saturated heterocyclic ring system optionally containing a further heteroatom selected from oxygen and nitrogen atoms, which ring system may be optionally substituted by one or more substituent groups independently selected from phenyl, -OR¹⁴, -COOR¹⁴, -NR¹⁵R¹⁶, -CONR¹⁵R¹⁶, -NR¹⁵COR¹⁶, -SONR¹⁵R¹⁶, NR¹⁵SO₂R¹⁶ or C₁-C₆ alkyl, itself

15 optionally substituted by one or more substituents independently selected from halogen atoms and -NR¹⁵R¹⁶ and -OR¹⁷ groups;

R¹⁰ represents a hydrogen atom or a C₁-C₆-alkyl or a phenyl group, the latter two of which may be optionally substituted by one or more substituent groups independently selected from halogen atoms, phenyl, -OR¹⁷ and -NR¹⁵R¹⁶; and

each of R⁷, R⁸, R⁹, R¹¹, R¹², R¹³, R¹⁴, R¹⁵, R¹⁶, R¹⁷ independently represents a hydrogen atom or a C₁-C₆, alkyl, or a phenyl group.

25 2. A compound according to claim 1, wherein R¹ represents an optionally substituted benzyl group.

30 3. A compound according to claim 1 or claim 2, wherein one of R² and R³ is hydrogen and the other is C₁-C₈ alkyl substituted by hydroxy and one or more methyl or ethyl groups.

4. A compound according to claim 1 selected from:

7-[(2-Hydroxy-1,1-dimethylethyl)amino]-5-[(phenylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

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(*R*)-7-[[1-(Hydroxymethyl)propyl]amino]-5-[(phenylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

(*R*)-7-[(2-Hydroxy-1-methylethyl)amino]-5-[(phenylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5 [[(2,3-Difluorophenyl)methyl]thio]-7-[(2-hydroxy-1,1-dimethylethyl)amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5 [[(2,3-Difluorophenyl)methyl]thio]-7-[(1*R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5 [[(2,3-difluorophenyl)methyl]thio]-7-[[2-(hydroxyethoxy)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

10 5 [[(2,3-difluorophenyl)methyl]thio]-7-[[2-hydroxy-1-(hydroxymethyl)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[(2-aminoethyl)amino]-5-[[2,3-difluorophenyl)methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

15 5 [[(2,3-difluorophenyl)methyl]thio]-7-[(2-hydroxyethyl)amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

N-[2-[[5-[[2,3-difluorophenyl)methyl]thio]-2,3-dihydro-2-oxothiazolo[4,5-*d*]pyrimidin-7-yl]amino]ethyl]methanesulfonamide,

18 (+/-)-5-[[2,3-difluorophenyl)methyl]thio]-7-[[2-(2-hydroxyethoxy)-1-

20 methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[[1*R*)-2-amino-1-methylethyl]amino]-5-[[2,3-difluorophenyl)methyl]thio] thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

25 5-[[2,3-difluorophenyl)methyl]thio]-7-[[1*R*)-2-[(2-hydroxyethyl)amino]-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[2,3-difluorophenyl)methyl]thio]-7-[[1*R*)-2-(dimethylamino)-1-

methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

30 5-[[4-(2-aminoethoxy)-3-chlorophenyl)methyl]thio]-7-[[1*R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[3-Chloro-4-methoxyphenyl)methyl]thio]-7-[[1*R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[3-Chloro-2-fluorophenyl)methyl]thio]-7-[(*(1R)*-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[2,3-Difluorophenyl)methyl]thio]-7-[(*(3R,4R)*-4-hydroxypyrrolidin-3-yl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[2,3-Difluorophenyl)methyl]thio]-7-[*(3R*)-pyrrolidin-3-ylamino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[[*(1R*)-2-Hydroxy-1-methylethyl]amino]-5-[[2-methyl-4-thiazolyl)methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[[2-Hydroxy-1-(hydroxymethyl)ethyl]amino]-5-[[2-methyl-4-thiazolyl)methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[(2-Hydroxy-1,1-dimethylethyl)amino]-5-[[2-methyl-4-thiazolyl)methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[(2-Hydroxy-1,1-dimethylethyl)amino]-5-[(2-methylphenyl)methyl]thio] thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

15 5-[(2-Furanylmethyl)thio]-7-[(*(1R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[[*(1R*)-2-Amino-1-methylethyl]amino]-5-[[3-chloro-2-fluorophenyl)methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one

(2*S*)-2-[[5-[[2,3-Difluorophenyl)methyl]thio]-2,3-dihydro-2-oxothiazolo[4,5-*d*]pyrimidin-*7*-yl]amino]-3-hydroxy-propanamide,

20 7-[[*(1R*)-2-hydroxy-1-methylethyl]amino]-5-[(2-thienylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

7-[[*(1R*)-2-hydroxy-1-methylethyl]amino]-5[[[3-methyl-4-(methylsulfonyl)phenyl]methyl]thio]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

25 5-[[[3-chloro-4-(trifluoromethoxy)phenyl)methyl]thio]-7-[[*(1R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[[2-fluoro-3-(trifluoromethyl)phenyl)methyl]thio]-7-[[*(1R*)-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[2,3-difluorophenyl)methyl]thio]-7-[2-[(dimethylamino)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

30 5-[[2,3-difluorophenyl)methyl]thio]-7-[2-[(dimethylamino)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(3*H*)-one,

5-[[(2-fluorophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

7-[[*(1R)*-2-hydroxy-1-methylethyl]amino]-5-[[2-methoxyphenyl)methyl]thio] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

5 7-[[*(1R)*-2-hydroxy-1-methylethyl]amino]-5-[(2-phenoxyethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

7-[[*(1R)*-2-hydroxy-1-methylethyl]amino]-5-[[3-methylphenyl)methyl]thio] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

10 5-[[2-fluoro-3-methylphenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

5-[[3-chlorophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

5-[[3-bromophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

15 5-[[[4-(difluoromethoxy)phenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

(+/-)-5-[[2,3-difluorophenyl)methyl]thio]-7-[[2-hydroxy-1-(methoxymethyl)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

7-[[2-hydroxy-1-(hydroxymethyl)ethyl]amino]-5-[(phenylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

20 5-[[2-bromophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

5-[[2-bromophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

5-[[2,3-Difluorophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

25 5-[[3-Chloro-2-fluorophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

(+/-)-5-[[2,3-difluorophenyl)methyl]thio]-7-[[2-hydroxy-1-(methoxymethyl)ethyl]amino]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

7-[[2-hydroxy-1-(hydroxymethyl)ethyl]amino]-5-[(phenylmethyl)thio]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

30 5-[[2,3-difluorophenyl)methyl]thio]-7-[[*(1R)*-2-hydroxy-1-methylethyl]amino] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,

7-[(*(1R)*-2-Hydroxy-1-methylethyl]amino]-5-[*(phenylmethyl)thio*]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,
5-[*(5-chloro-1,2,3-thiadiazol-4-yl)thio*]-7-[(*(1R)*-2-hydroxy-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,
and their pharmaceutically acceptable salts and solvates.

5. A compound according to claim 1 selected from:

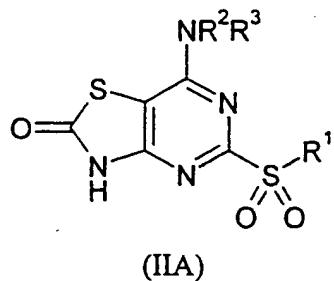
5-[(2,3-Difluorophenyl)methyl]thio]-7-[(*(1R)*-2-hydroxy-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one sodium salt,
10 5-[(3-Chloro-2-fluorophenyl)methyl]thio]-7-[(*(1R)*-2-hydroxy-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one sodium salt,
(+/-)-5-[(2,3-difluorophenyl)methyl]thio]-7-[(2-hydroxy-1-(methoxymethyl)ethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one sodium salt,
15 7-[(2-hydroxy-1-(hydroxymethyl)ethyl]amino]-5-[*(phenylmethyl)thio*]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one sodium salt, or
7-[(*(1R)*-2-Hydroxy-1-methylethyl]amino]-5-[*(phenylmethyl)thio*]thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one sodium salt.

6. A compound according to claim 1 selected from:

20 7-[(*(1R)*-2-amino-1-methylethyl]amino]-5-[(2,3-difluorophenyl)methyl]thio] thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one trifluoroacetate,
5-[(2,3-difluorophenyl)methyl]thio]-7-[(*(1R)*-2-[(2-hydroxyethyl)amino]-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one trifluoroacetate,
25 5-[(2,3-difluorophenyl)methyl]thio]-7-[(*(1R)*-2-(dimethylamino)-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one,
5-[[4-(2-aminoethoxy)-3-chlorophenyl]methyl]thio]-7-[(*(1R)*-2-hydroxy-1-methylethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one trifluoroacetate,
30 5-[(2,3-difluorophenyl)methyl]thio]-7-[2-[(dimethylamino)ethyl]amino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one monohydrochloride, or
5-[(2,3-Difluorophenyl)methyl]thio]-7-[(3*R*)-pyrrolidin-3-ylamino]-thiazolo[4,5-*d*]pyrimidin-2(*3H*)-one dihydrochloride.

7. A process for the preparation of a compound of formula (I) as defined in claim 1 which comprises:

(a) treating a compound of formula (IIA):

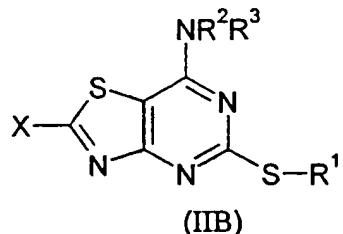


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where R¹, R² and R³ are as defined in formula (I) with a thiol R¹SH in the presence of a suitable base, or

10

(b) treatment of a compound of formula (IIB):



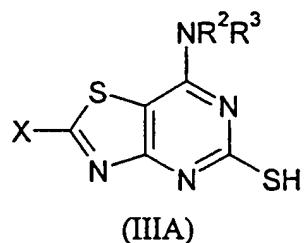
15

where R¹, R² and R³ are as defined in formula (I) and X is a leaving group with a metal alkoxide, followed by treatment with an acid or base, and optionally after (a) or (b) forming a pharmaceutically acceptable salt.

20

8. A compound of formula (IIA) or (IIB) as defined in claim 7.

9. A compound of formula (IIIA):



25

where R² and R³ are as defined in formula (I) and X is NH₂

PA 2 10 10. A pharmaceutical composition comprising a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6 in association with a pharmaceutically acceptable adjuvant, diluent or carrier.

11. A process for the preparation of a pharmaceutical composition as claimed in claim 10 which comprises mixing a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6 with a pharmaceutically acceptable adjuvant, diluent or carrier.

12. A compound of formula (I), or a pharmaceutically-acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6, for use in therapy.

15 13. Use of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6 in the manufacture of a medicament for use in therapy.

20 14. A method of treating a chemokine mediated disease wherein the chemokine binds to one or more chemokine receptors, which comprises administering to a patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6.

25 15. A method according to claim 14 in which the chemokine receptor belongs to the CXC chemokine receptor subfamily.

PA 3 30 16. A method according to claim 14 or 15 in which the chemokine receptor is the CXCR2 receptor.

35 17. A method of treating an inflammatory disease in a patient suffering from, or at risk of, said disease, which comprises administering to the patient a therapeutically effective amount of a compound of formula (I), or a pharmaceutically acceptable salt or solvate thereof, as claimed in any one of claims 1 to 6.

18. A method according to claim 17, wherein the disease is psoriasis, a disease in which angiogenesis is associated with raised CXCR2 chemokine levels, or COPD.

19. A method according to claim 15, wherein the disease is psoriasis.